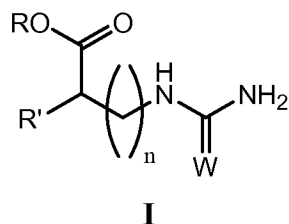


**IN THE CLAIMS**

Please replace all previous listings, and versions, of the claims with the following claims, where added text is indicated by underlining and deleted text is indicated by strikethrough:

1. (Original) A compound, comprising a non-protein-binding moiety (NPBM) and at least one protein-binding group (PBG).
2. (Original) The compound of claim 1, wherein the NPBM is a polyol, sugar, amino acid, or dendrimer moiety.
3. (Original) The compound of claim 1, wherein the NPBM is a polyol moiety; and said polyol moiety is a sorbitol or mannitol moiety.
4. (Original) The compound of claim 1, wherein the NPBM is a sugar moiety; and said sugar moiety is a glucose, sucrose, or trehalose moiety.
5. (Original) The compound of claim 1, wherein the NPBM is an amino acid moiety; and said amino acid moiety is an arginine betaine, proline, or ectoine moiety.
6. (Original) The compound of claim 1, wherein the NPBM is a dendrimer moiety; and said dendrimer moiety is based on benzene, pentaerythritol,  $P(CH_2OH)_3$ , or TRIS.
7. (Original) The compound of any of claims 1-6, wherein the PBG is a urea, guanidinium ion, detergent, amino acid, denaturant, surfactant, polysorbate, polaxamer, citrate, chaotrope, or acetate group.
8. (Original) The compound of any of claims 1-6, wherein the PBG is a guanidinium ion.
9. (Original) The compound of any of claims 1-6, wherein the PBG is sodium dodecyl sulfate.

10. (Original) A compound represented by formula **I**:



wherein:

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal;

R' is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or (R'')<sub>3</sub>N;

R'' is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, or heteroaralkyl;

W is O, NH<sub>2</sub><sup>+</sup>(halogen)<sup>-</sup>, or S; and

n is 1, 2, or 4-100.

11. (Original) The compound of claim 10, wherein R is an electron pair.
12. (Original) The compound of claim 10, wherein R' is H.
13. (Original) The compound of claim 10, wherein R' is (R'')<sub>3</sub>N.
14. (Original) The compound of claim 10, wherein R' is H<sub>3</sub>N<sup>+</sup>.
15. (Original) The compound of claim 10, wherein W is NH<sub>2</sub><sup>+</sup>Cl<sup>-</sup>.
16. (Original) The compound of claim 10, wherein n is 1.
17. (Original) The compound of claim 10, wherein n is 2.
18. (Original) The compound of claim 10, wherein n is 4.
19. (Original) The compound of claim 10, wherein n is 5.

20. (Original) The compound of claim 10, wherein n is 6.
21. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 1.
22. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 2.
23. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 4.
24. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 5.
25. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 6.
26. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is O, and n is 1.
27. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is O, and n is 2.
28. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is O, and n is 4.
29. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is O, and n is 5.

30. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}_3\text{N}^+$ , W is O, and n is 6.

31. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 1.

32. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 2.

33. (Original) The compound of claim 10, wherein R is an electron pair, R' is  $\text{H}^+$ , W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 4.

34. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 5.

35. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is  $\text{NH}_2^+\text{Cl}^-$ , and n is 6.

36. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 1.

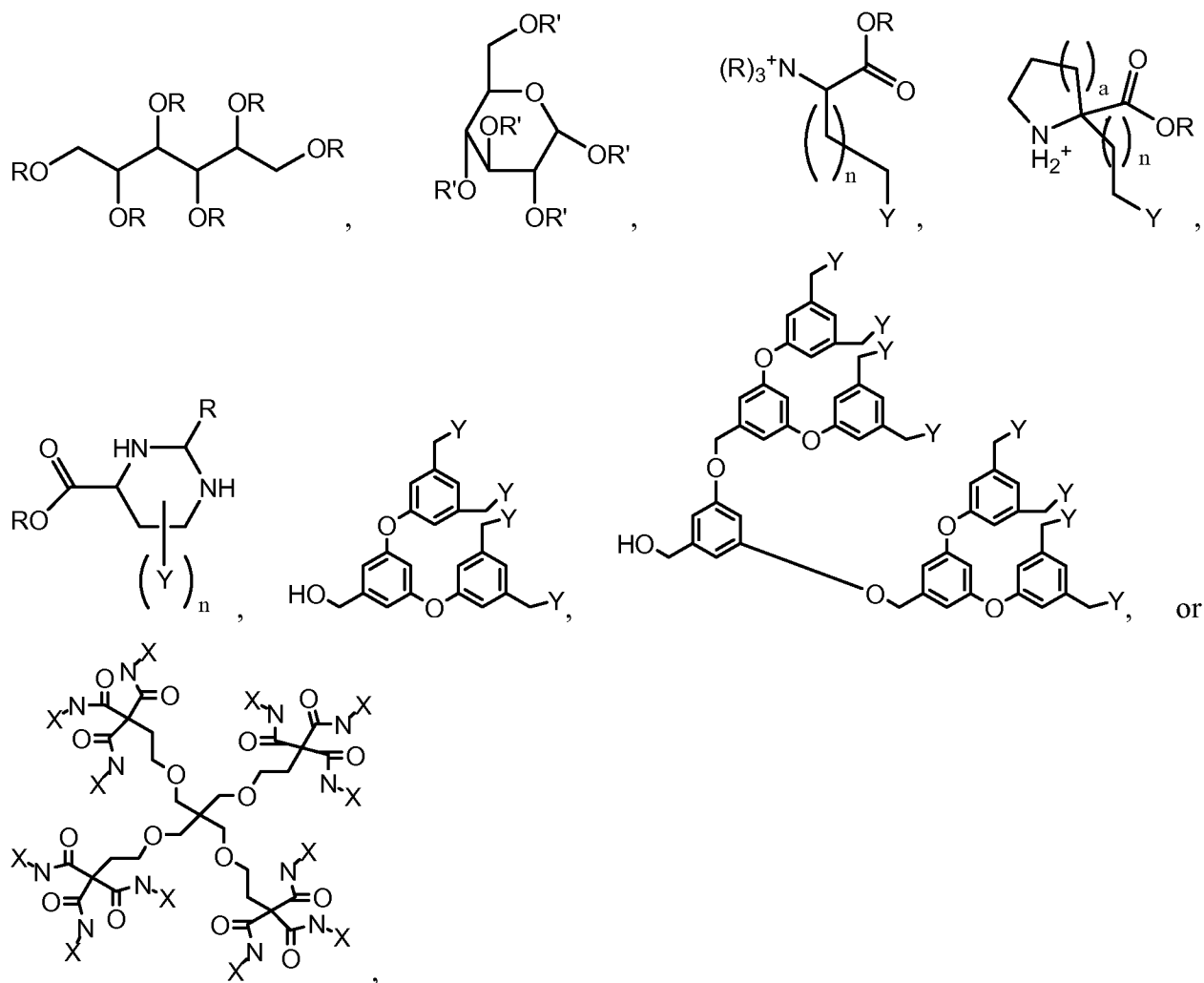
37. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 2.

38. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 4.

39. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 5.

40. (Original) The compound of claim 10, wherein R is an electron pair, R' is H, W is O, and n is 6.

41. (Original) A compound selected from the group consisting of:



wherein, independently for each occurrence,

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, an alkali metal, or CH<sub>2</sub>Y;

R' is H, a sugar radical, or CH<sub>2</sub>Y;

n is an integer from 1 to 100, inclusive;

a is 1, 2, or 3;

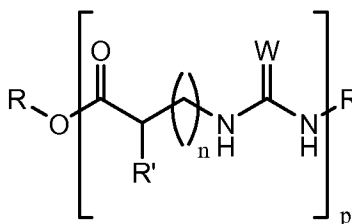
X is C(CH<sub>2</sub>Y)<sub>3</sub>; and

Y is a protein binding group,

wherein at least one Y is present in all compounds.

42. (Original) The compound of claim 41, wherein Y is a guanidinium ion.

43. (Original) A polymer of formula **II**, **III**, **IV**, **V**, **VI**, **VII**, **VIII**, or **IX**:



**II**

wherein, independently for each occurrence:

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal;

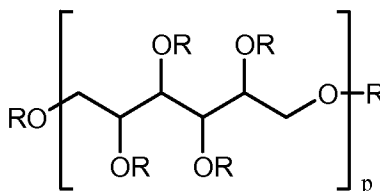
R' is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or (R'')<sub>3</sub>N;

R'' is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, or heteroaralkyl;

W is O, NH<sub>2</sub><sup>+</sup>(halogen)<sup>-</sup>, or S;

n is 1, 2, or 4-100; and

p is an integer from 2 to 1000 inclusive;



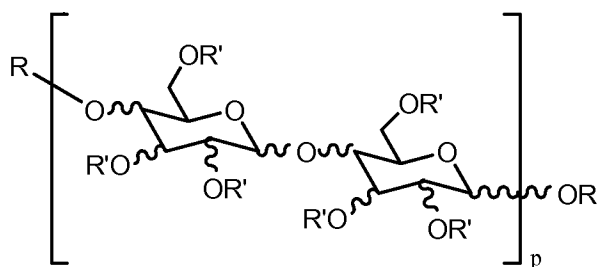
**III**

wherein, independently for each occurrence,

R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal, or CH<sub>2</sub>Y;

p is an integer from 2 to 1000 inclusive; and

Y is a PBG, wherein at least one Y is present;

**IV**

wherein, independently for each occurrence:

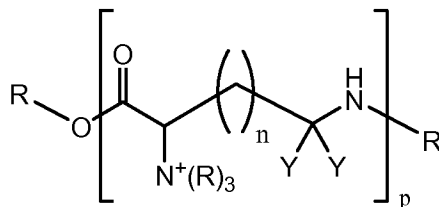
R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal, or  $\text{CH}_2\text{Y}$ ;

R' is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or  $(\text{R}'')_3\text{N}$ ;

R'' is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, or heteroaralkyl;

p is an integer from 2 to 1000 inclusive; and

Y is a PBG, wherein at least one Y is present;

**V**

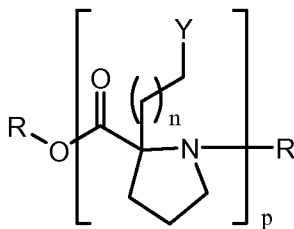
wherein, independently for each occurrence:

R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal, or  $\text{CH}_2\text{Y}$ ;

n is an integer from 1 to 100 inclusive;

p is an integer from 2 to 1000 inclusive; and

Y is a PBG;

**VI**

wherein, independently for each occurrence,

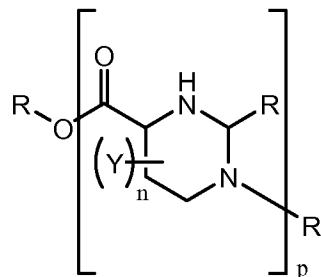
R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, an alkali metal, or  $\text{CH}_2\text{Y}$ ;

n is an integer from 1 to 100, inclusive;

a is 1, 2, or 3;

Y is a PBG; and

p is an integer from 2 to 1000, inclusive;



**VII**

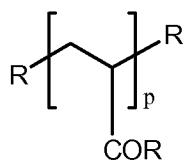
wherein, independently for each occurrence,

R is H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, an alkali metal, or CH<sub>2</sub>Y;

n is an integer from 1 to 6, inclusive;

Y is a PBG; and

p is an integer from 2 to 1000, inclusive; or

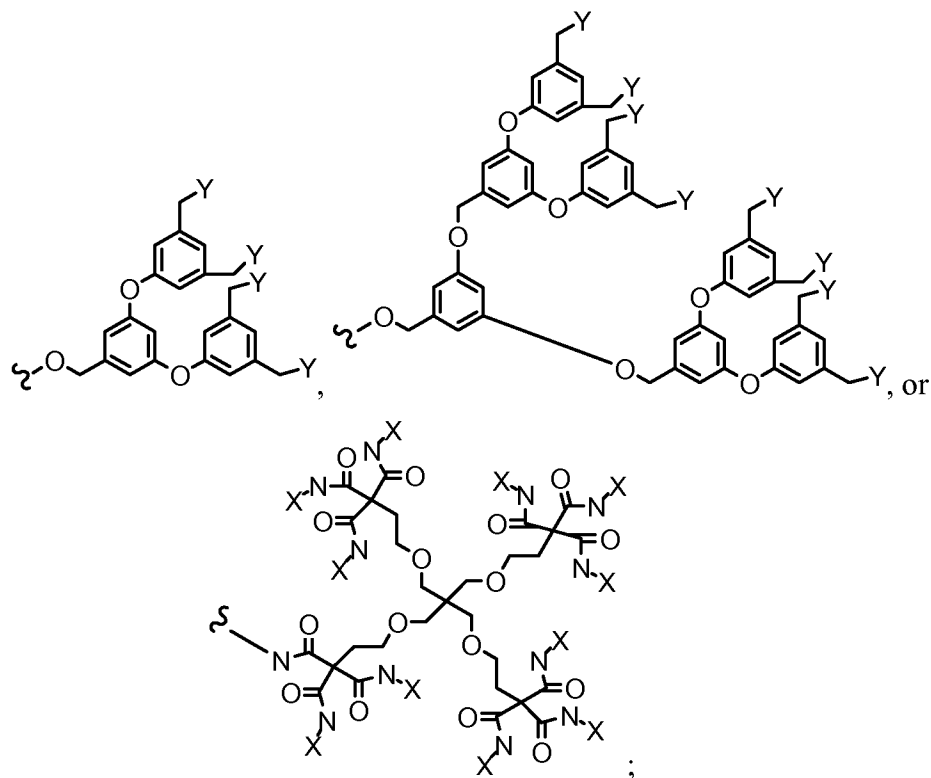


**VIII**

wherein, independently for each occurrence,

R is H, OH, alkyl, alkoxy, aryl, heteroaryl, aralkyl, heteroaralkyl, -O-alkali metal, CH<sub>2</sub>Y, OCH<sub>2</sub>Y, or has a structure selected from the following:



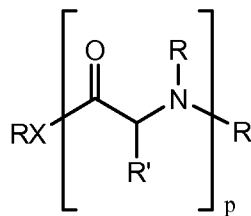


a is 1, 2, or 3;

X is C(CH<sub>2</sub>Y)<sub>3</sub>;

Y is a PBG, wherein at least one Y is present; and

p is an integer from 2 to 1000, inclusive; or



## IX

wherein, individually for each occurrence:

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal;

R' is a sidechain of an alpha-amino acid, wherein at least one instance of R' is the sidechain of arginine;

X is O or NR; and

p is an integer from 2 to 1000, inclusive.

44. (Original) A method of screening compounds or polymers for the property of inhibiting protein aggregation in solution, comprising:

- a) computing a set of parameters utilizing molecular modeling based on compounds or polymers known to have the property of inhibiting protein aggregation;
- b) applying those parameters to other compounds or polymers; and
- c) choosing the compounds or polymers that meet the criteria of those parameters.

45. (Original) A method of preparing a compound or polymers having the property of protein aggregation inhibition in solution, comprising:

- a) computing a set of parameters utilizing molecular modeling based on compounds or polymers known to have the property of inhibiting protein aggregation;
- b) designing a compound or polymer having the property of protein aggregation inhibition in solution based on those parameters; and
- c) synthesizing the compound or polymer having the property of protein aggregation inhibition in solution.

46. (Original) A method of classifying a compound or polymer as either inhibitory of protein aggregation in solution or not inhibitory of protein aggregation in solution, comprising:

- a) computing a set of parameters utilizing molecular modeling based on compounds or polymers known to have the property of inhibiting protein aggregation;
- b) applying those parameters to a compound or polymer; and
- c) classifying the compound or polymer that meet the criteria of those parameters as inhibitory of protein aggregation in solution.

47. (Original) A method of determining the preferential binding coefficient,  $\Gamma_{XP}$ , of an additive in a protein solution, comprising:

- a) determining the phase space trajectories of the protein, solvent, and additive using molecular dynamics;
- b) calculating the distance,  $r$ , between the center of mass for both the solvent molecule and additive molecule to the protein's van der Waals surface;

c) determining the minimum distance,  $r^*$ , at which no significant differences between the local ( $r = r^*$ ) and bulk density are observed;

d) determining which molecules lie within the distance,  $r^*$ , from the protein surface and classifying these molecules as the local domain;

e) determining which molecules lie outside the distance,  $r^*$ , from the protein surface and classifying these molecules as the bulk domain;

f) determining the instantaneous preferential binding coefficient,  $\Gamma_{XP}(t)$ , using the following formula:

$$\Gamma_{XP}(t) = n_X^{\text{II}} - n_X^{\text{I}} (n_W^{\text{II}} / n_W^{\text{I}})$$

wherein:

$n_X^{\text{II}}$  = the number of additive molecules in the bulk domain;

$n_X^{\text{I}}$  = the number of additive molecules in the local domain;

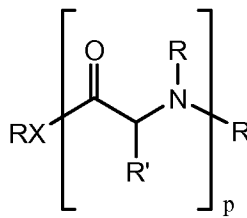
$n_W^{\text{II}}$  = the number of solvent molecules in the bulk domain; and

$n_W^{\text{I}}$  = the number of solvent molecules in the local domain; and

g) calculating the preferential binding coefficient,  $\Gamma_{XP}$ , as the time average of each of the values in step f) using the following formula:

$$\Gamma_{XP} = \frac{1}{t} \int_0^t \Gamma_{XP}(t') dt'.$$

48. (Currently amended) A method of suppressing or preventing aggregation of a protein in solution, comprising the step of combining in a solution ~~the compound or~~ (i) a polymer of formula IX ~~any of claims 1 to 43~~



**IX**

wherein, individually for each occurrence:

R is an electron pair, H, alkyl, aryl, heteroaryl, aralkyl, heteroaralkyl, or an alkali metal;

R' is a sidechain of an alpha-amino acid, wherein at least one instance of R' is the sidechain of arginine;

X is O or NR; and

p is an integer from 2 to 1000, inclusive; and

(ii) a protein.

49. (Original) The method of claim 48, wherein the protein is a recombinant protein.
50. (Original) The method of claim 48, wherein the protein is a recombinant antibody.
51. (Original) The method of claim 48, wherein the protein is a recombinant human antibody.
52. (Original) The method of claim 48, wherein the protein is a recombinant human protein.
53. (Original) The method of claim 48, wherein the protein is recombinant human insulin, recombinant human erythropoietin or a recombinant human interferon.
54. (Original) The method of claim 48, wherein the solution is an aqueous solution.
55. (Original) The method of claim 48, wherein the protein is a recombinant protein; and the solution is an aqueous solution.
56. (Original) The method of claim 48, wherein the protein is a recombinant human protein; and the solution is an aqueous solution.
57. (Original) A method of decreasing the toxicological risk associated with administering a protein to a mammal in need thereof, comprising the steps of adding to a first solution of a protein a compound or polymer of any of claims 1 to 43 to give a second solution; and administering to a mammal in need thereof a therapeutic amount of said second solution.

- 58. (Original) The method of claim 57, wherein the protein is a recombinant protein.
- 59. (Original) The method of claim 57, wherein the protein is a recombinant antibody.
- 60. (Original) The method of claim 57, wherein the protein is a recombinant human antibody.
- 61. (Original) The method of claim 57, wherein the protein is a recombinant mammalian protein.
- 62. (Original) The method of claim 57, wherein the protein is a recombinant human protein.
- 63. (Original) The method of claim 57, wherein the protein is recombinant human insulin, recombinant human erythropoietin or a recombinant human interferon.
- 64. (Original) The method of claim 57, wherein the first solution and the second solution are aqueous solutions.
- 65. (Original) The method of claim 57, wherein the protein is a recombinant protein; and the first solution and the second solution are aqueous solutions.
- 66. (Original) The method of claim 57, wherein the protein is a recombinant human antibody; and the first solution and the second solution are aqueous solutions.
- 67. (Original) The method of claim 57, wherein the protein is a recombinant human protein; and the first solution and the second solution are aqueous solutions.

68. (Original) A method of facilitating native folding of a recombinant protein in solution, comprising the step of combining in a solution a compound or polymer of any of claims 1 to 43 and a recombinant protein.
69. (Original) The method of claim 68, wherein the recombinant protein is a recombinant antibody.
70. (Original) The method of claim 68, wherein the recombinant protein is a recombinant human antibody.
71. (Original) The method of claim 68, wherein the recombinant protein is a recombinant mammalian protein.
72. (Original) The method of claim 68, wherein the recombinant protein is a recombinant human protein.
73. (Original) The method of claim 68, wherein the recombinant protein is recombinant human insulin, recombinant human erythropoietin or a recombinant human interferon.
74. (Original) The method of claim 68, wherein the solution is an aqueous solution.
75. (Original) The method of claim 68, wherein the recombinant protein is a recombinant human antibody; and the solution is an aqueous solution.
76. (Original) The method of claim 68, wherein the recombinant protein is a recombinant human protein; and the solution is an aqueous solution.